We claim:

- 1. A method for inhibiting an altered growth state of a cell having a ptc loss-of-function phenotype or a smoothened gain-of-function phenotype, comprising contacting the cell with a ptc agonist in a sufficient amount to inhibit the altered growth state, wherein the ptc agonist is a organic molecule having a molecular weight less than about 750 amu.
- 2. A method for inhibiting aberrant proliferation of a cell having a ptc loss-of-function phenotype or a smoothened gain-of-function phenotype comprising contacting the cell with a ptc agonist in a sufficient amount to inhibit proliferation of the cell.
- 3. The method of claim 1, wherein the *ptc* agonist causes repression of *smoothened*-mediated signal transduction.
- 4. The method of claim 1, wherein the ptc agonist is a steroidal alkaloid.
- 5. The method of claim 4, wherein the steroidal alkaloid is represented in the general forumlas (I), or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

Formula I

wherein, as valence and stability permit,

R₂, R₃, R₄, and R₅, represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈;

 R_6 , R_7 , and R_7 , are absent or represent, independently, halogens, alkyls, alkenyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH_2)_m- R_8 , or

R₆ and R₇, or R₇ and R'₇, taken together form a ring or polycyclic ring, e.g., which is susbstituted or unsubstituted,

with the proviso that at least one of R₆, R₇, or R'₇ is present and includes a primary or secondary amine;

Rg represents an aryl, a cycloalkyl, a cycloalkenyl, a heterocycle, or a polycycle; and

m is an integer in the range 0 to 8 inclusive.

6. The method of claim 5, wherein:

 R_2 and R_3 , for each occurrence, is an -OH, alkyl, -O-alkyl, -C(O)-alkyl, or -C(O)- R_8 ;

 R_4 , for each occurrence, is an absent, or represents -OH, =O, alkyl, -O-alkyl, -C(O)-alkyl, or -C(O)- R_8 ;

 R_6 , R_7 , and R'_7 each independently represent, hydrogen, alkyls, alkenyls, alkynyls, amines, imines, amides, carbonyls, carboxyls, carboxamides, ethers, thioethers, esters, or -(CH₂)_m-R₈, or

R₇, and R'₇ taken together form a furanopiperidine, such as perhydrofuro[3,2-b]pyridine, a pyranopiperidine, a quinoline, an indole, a pyranopyrrole, a naphthyridine, a thiofuranopiperidine, or a thiopyranopiperidine

with the proviso that at least one of R_6 , R_7 , or R'_7 is present and includes a primary or secondary amine;

 R_8 represents an aryl, a cycloalkyl, a cycloalkenyl, a heterocycle, or a polycycle, and preferably R_8 is a piperidine, pyrimidine, morpholine, thiomorpholine, pyridazine,

7. The method of claim 4, wherein the steroidal alkaloid is represented in the general formula (II), or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

Formula II

wherein

 R_2 , R_3 , R_4 , and R_5 , represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈;

R₆, R₇, and R'₇, are absent or represent, independently, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol,

amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈, or

R₆ and R₇, or R₇ and R'₇, taken together form a ring or polycyclic ring, e.g., which is susbstituted or unsubstituted,

with the proviso that at least one of R_6 , R_7 , or R_7 is present and includes a primary or secondary amine;

X represents O or S, though preferably O.

8. The method of claim 4, wherein the steroidal alkaloid is represented in the general formula (III), or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

$$R3$$
 $R4$
 $R5$
 $R9$
 $R3$
 $R3$
 $R4$
 $R5$
 $R5$
 $R9$
 $R3$
 $R1$
 $R9$
 $R2$
 $R3$
 $R4$
 $R5$
 $R5$
 $R9$
 $R9$

Formula III

wherein

 R_2 , R_3 , R_4 , and R_5 , represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈;

Rg represents an aryl, a cycloalkyl, a cycloalkenyl, a heterocycle, or a polycycle; and

A and B represent monocyclic or polycyclic groups;

T represent an alkyl, an aminoalkyl, a carboxyl, an ester, an amide, ether or amine linkage of 1-10 bond lengths;

T' is absent, or represents an alkyl, an aminoalkyl, a carboxyl, an ester, an amide, ether or amine linkage of 1-3 bond lengths, wherein if T and T' are present together, than T and T' taken together with the ring A or B form a covelently closed ring of 5-8 ring atoms;

R9 represent one or more substitutions to the ring A or B, which for each occurrence, independently represent halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or $-(CH_2)_m$ -R8; and

n and m are, independently, zero, 1 or 2;

with the proviso that A and R₉, or T, T' B and R₉, taken together include at least one primary or secondary amine.

9. The method of claim 4, wherein the steroidal alkaloid is represented in the general formula (IV), or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

Formula IV

wherein

R₂, R₃, R₄, and R₅, represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy,

amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or $-(CH_2)_m-R_8$;

 R_6 is absent or represent, independently, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈;

R9 represent one or more substitutions to the ring A or B, which for each occurrence, independently represent halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or $-(CH_2)_m$ -R8; and

R₂₂ is absent or represents an alkyl, an alkoxyl or -OH.

10. The method of claim 4, wherein the steroidal alkaloid is represented in the general formula (V) or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

$$R4$$
 $R6$
 $R9$
 $R4$
 $R6$
 $R9$
 $R4$
 $R6$
 $R9$
 $R9$
 $R1$
 $R9$
 $R2$
 $R1$
 $R2$
 $R2$
 $R3$
 $R4$
 $R6$
 $R9$
 $R9$

Formula V

R3

or

R2

 R_2 , R_3 , R_4 , and R_5 , represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈;

R₆ is absent or represent, independently, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈; and

R9 represent one or more substitutions to the ring A or B, which for each occurrence, independently represent halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carboxyls, carboxyls, carboxamides,

anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or $-(CH_2)_m-R_8$.

11. The method of claim 4, wherein the steroidal alkaloid is represented in the general formula (VI), or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

$$R4$$
 $R9$
 $R4$
 $R9$
 $R4$
 $R9$
 $R4$
 $R9$
 $R9$
 $R1$
 $R2$
 $R2$
 $R3$
 $R4$
 $R5$
 $R5$

Formula VI

wherein

R₂, R₃, R₄, and R₅, represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈; and

R9 represent one or more substitutions to the ring A or B, which for each occurrence, independently represent halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or $-(CH_2)_m$ -R₈.

12. The method of claim 4, wherein the steroidal alkaloid is represented in the general formula (VII) or unsaturated forms thereof and/or seco-, nor- or homo-derivatives thereof:

Formula VII

wherein

R₂, R₃, R₄, and R₅, represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =O, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈; and

R9 represent one or more substitutions to the ring A or B, which for each occurrence, independently represent halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or $-(CH_2)_m$ -R₈.

13. The method of claim 4, wherein the steroidal alkaloid does not substantially interfere with the biological activity of such steroids as aldosterone, androstane, androstene, androstenedione, androsterone, cholecalciferol, cholestane, cholic acid, corticosterone, cortisol, cortisol acetate, cortisone, cortisone acetate, deoxycorticosterone, digitoxigenin, ergocalciferol, ergosterol, estradiol-17-α, estradiol-17-β, estriol, estrane, estrone, hydrocortisone, lanosterol, lithocholic

acid, mestranol, β -methasone, prednisone, pregnane, pregnenolone, progesterone, spironolactone, testosterone, triamcinolone and their derivatives.

- 14. The method of claim 4, wherein the steroidal alkaloid does not specifically bind a nuclear hormone receptor.
- 15. The method of claim 4, wherein the steroidal alkaloid does not specifically bind estrogen or testerone receptors.
- 16. The method of claim 4, wherein the steroidal alkaloid has no estrogenic activity at therapeutic concentrations.
- 17. The method of claim 1, wherein the *ptc* agonist inhibits *ptc* loss-of-function or *smoothened* gain-of-function mediated signal transduction with an ED₅₀ of 1mM or less.
- 18. The method of claim 1, wherein the ptc agonist inhibits ptc loss-of-function or smoothened gain-of-function mediated signal transduction with an ED₅₀ of 1μM or less.
- 19. The method of claim 1, wherein the *ptc* agonist inhibits *ptc* loss-of-function or *smoothened* gain-of-function mediated signal transduction with an ED₅₀ of 1nM or less.
- 20. The method of claim 1, wherein the cell is contacted with the ptc agonist in vitro.
- 21. The method of claim 1, wherein the cell is contacted with the ptc agonist in vivo.
- 22. The method of claim 1, wherein the *ptc* agonist is administered as part of a therapeutic or cosmetic application.

- 23. The method of claim 22, wherein the therapeutic or cosmetic application is selected from the group consisting of regulation of neural tissues, bone and cartilage formation and repair, regulation of spermatogenesis, regulation of smooth muscle, regulation of lung, liver and other organs arising from the primative gut, regulation of hematopoietic function, regulation of skin and hair growth, etc.
- 24. A pharmaceutical preparation comprising a steroidal alkaloid represented in the general forumlas (I), or unsaturated forms thereof and/or seco-, nor- or homoderivatives thereof:

Formula I

wherein, as valence and stability permit,

 R_2 , R_3 , R_4 , and R_5 , represent one or more substitutions to the ring to which each is attached, for each occurrence, independently represent hydrogen, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carboxyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH₂)_m-R₈;

 R_6 , R_7 , and R_7 , are absent or represent, independently, halogens, alkyls, alkenyls, alkynyls, aryls, hydroxyl, =0, =S, alkoxyl, silyloxy, amino, nitro, thiol, amines, imines, amides, phosphoryls, phosphonates, phosphines, carbonyls, carboxyls, carboxamides, anhydrides, silyls, ethers, thioethers, alkylsulfonyls, arylsulfonyls, selenoethers, ketones, aldehydes, esters, or -(CH_2)_m- R_8 , or

R₆ and R₇, or R₇ and R'₇, taken together form a ring or polycyclic ring, e.g., which is susbstituted or unsubstituted,

with the proviso that at least one of R₆, R₇, or R'₇ is present and includes a primary or secondary amine;

Rg represents an aryl, a cycloalkyl, a cycloalkenyl, a heterocycle, or a polycycle; and

m is an integer in the range 0 to 8 inclusive.

- 25. A method for inhibiting an altered growth state of a cell having a ptc loss-of-function phenotype, hedgehog gain-of-function phenotype, or a smoothened gain-of-function phenotype, comprising contacting the cell with a composition including at least one cAMP agonist.
- 26. The method of claim 25, wherein at least one cAMP agonist activates adenylate cyclase.
- 27. The method of claim 25, wherein at least one cAMP agonist is a cAMP analog.
- 28. The method of claim 25, wherein at least one cAMP agonist is a cAMP phosphodiesterase inhibitor.
- 29. The method of claim 25, wherein the composition inhibits ptc loss-of-function, hedgehog gain-of-function, or smoothened gain-of-function mediated signal transduction with an ED₅₀ of 1 mM or less.
- 30. The method of claim 25, wherein the composition inhibits *ptc* loss-of-function, hedgehog gain-of-function, or smoothened gain-of-function mediated signal transduction with an ED₅₀ of 1 μM or less.
- 31. The method of claim 25, wherein the composition inhibits ptc loss-of-function, hedgehog gain-of-function, or smoothened gain-of-function mediated signal transduction with an ED₅₀ of 1 nM or less.

- 32. The method of claim 25, wherein the cell is contacted with the composition in vitro.
- 33. The method of claim 25, wherein the cell is contacted with the composition in vivo.
- 34. The method of claim 25, wherein the composition is administered as part of a therapeutic or cosmetic application.
- 35. The method of claim 34, wherein the therapeutic or cosmetic application is selected from the group consisting of regulation of neural tissues, bone and cartilage formation and repair, regulation of spermatogenesis, regulation of smooth muscle, regulation of lung, liver and other organs arising from the primative gut, regulation of hematopoietic function, regulation of skin and hair growth, etc.
- 36. The method of claim 25, wherein the composition includes forskolin or a derivative thereof.
- 37. A method for treating or preventing basal cell carcinoma, comprising administering a composition including a cAMP agonist to a patient in an amount sufficient to inhibit progression of basal cell carcinoma.
- 38. A method for inhibiting an altered growth state of a cell having a ptc loss-of-function phenotype, hedgehog gain-of-function phenotype, or a smoothened gain-of-function phenotype, comprising

determining the phenotype of the cell; and

if the phenotype is a ptc loss-of-function, hedgehog gain-of-function, or a smoothened gain-of-function phenotype, treating the cell with a cAMP agonist in an amount sufficient to inhibit the altered growth state of the cell.